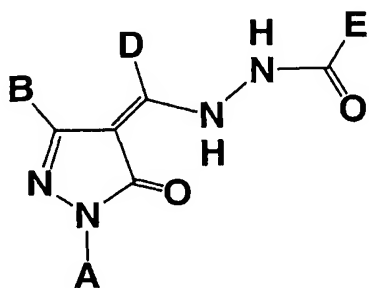


AMENDMENTS TO THE SPECIFICATION

Please amend the paragraph beginning on page 7, line 5 as follows:

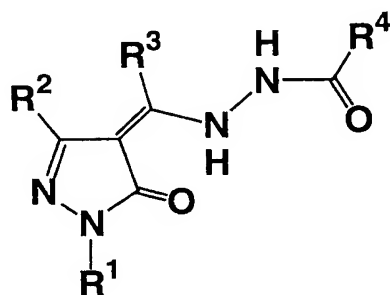
Namely, the present invention relates to a pyrazolone compound represented by the formula (1)



Formula (1)

wherein A is a C<sub>2-14</sub> aryl group (the C<sub>2-14</sub> aryl group may be optionally substituted with one or more C<sub>1-6</sub> alkyl groups, one or more C<sub>1-3</sub> alkyl groups substituted with one or more fluorine atoms, one or more halogen atoms, one or more nitro groups, one or more C<sub>1-6</sub> alkylcarbonyl groups, one or more hydroxyl groups or one or more amino groups (the hydroxyl group and the amino group may be substituted with a C<sub>1-6</sub> alkyl group or a C<sub>1-6</sub> alkylcarbonyl group)), B is a hydrogen atom, a C<sub>1-6</sub> alkyl group, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms or a C<sub>2-14</sub> aryl group, D is a hydrogen atom, a C<sub>1-6</sub> alkyl group, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms or a C<sub>2-14</sub> aryl group, and E is a C<sub>2-14</sub> aryl group (the C<sub>2-14</sub> aryl group is optionally substituted with one or more hydroxyl groups, one or more nitro groups, one or more halogen atoms, one or more cyano groups, one or more C<sub>1-3</sub> alkyl groups substituted with one or more fluorine atoms, NG<sup>1</sup>G<sup>2</sup> (wherein G<sup>1</sup> and G<sup>2</sup> are independently hydrogen atoms, formyl groups, C<sub>1-6</sub> alkyl groups or C<sub>1-6</sub> alkylcarbonyl groups), one or more carboxyl groups, one or more sulfonic acid groups, one or more phosphonic acid groups, one or more carbamido ~~earbamoyl~~ groups (the carbamido ~~earbamoyl~~

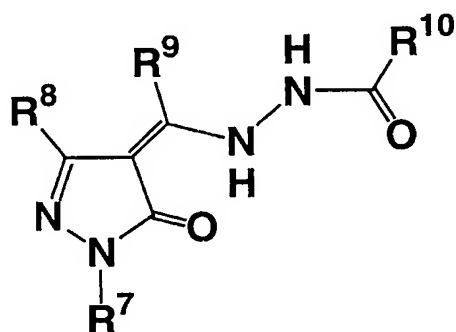
group may be substituted with a C<sub>1-6</sub> alkyl group), one or more sulfamido ~~sulfamoyl~~ groups (the sulfamido ~~sulfamoyl~~ group may be substituted with a C<sub>1-6</sub> alkyl group), one or more hydroxycarbamido ~~hydroxycarbameoyl~~ groups, one or more hydroxysulfamido ~~hydroxysulfameoyl~~ groups, one or more tetrazole groups, one or more C<sub>1-6</sub> alkoxy carbonyl groups or X(CYZ)<sub>n</sub>CO<sub>2</sub>H (wherein X is CH<sub>2</sub>, O, S or NG<sup>3</sup> (G<sup>3</sup> is a hydrogen atom, a C<sub>1-6</sub> alkyl group, a formyl group or a C<sub>1-6</sub> alkyl carbonyl group), Y and Z are independently hydrogen atoms or C<sub>1-3</sub> alkyl groups, and n is 0, 1, 2 or 3)), a tautomer, prodrug or pharmaceutically acceptable salt of the compound or a solvate thereof, a thrombopoietin receptor activator, a preventive, therapeutic or improving agent for diseases against which activation of the thrombopoietin receptor is effective which contains the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient, and a platelet increasing agent containing the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient. It also relates to a pyrazolone compound represented by the formula (2)



Formula (2)

wherein R<sup>1</sup> is a C<sub>2-14</sub> aryl group (the C<sub>2-14</sub> aryl group may be optionally substituted with one or more C<sub>1-6</sub> alkyl groups, one or more C<sub>1-3</sub> alkyl groups substituted with one or more

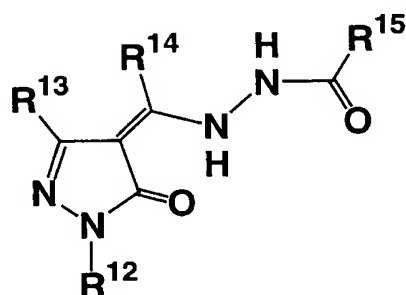
fluorine atoms, one or more halogen atoms, one or more nitro groups, one or more C<sub>1-6</sub> alkylcarbonyl groups, one or more hydroxyl groups or one or more amino groups (the hydroxyl group and the amino group may be substituted with a C<sub>1-6</sub> alkyl group or a C<sub>1-6</sub> alkylcarbonyl group)), R<sup>2</sup> is a hydrogen atom, a C<sub>1-6</sub> alkyl group, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms or a C<sub>2-14</sub> aryl group, R<sup>3</sup> is a hydrogen atom, a C<sub>1-6</sub> alkyl group, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms or a C<sub>2-14</sub> aryl group, and R<sup>4</sup> is a C<sub>2-14</sub> aryl group (the C<sub>2-14</sub> aryl group is optionally substituted with one or more hydroxyl groups, one or more nitro groups or NR<sup>5</sup>R<sup>6</sup> (wherein R<sup>5</sup> and R<sup>6</sup> are independently hydrogen atoms, formyl groups, C<sub>1-6</sub> alkyl groups or C<sub>1-6</sub> alkylcarbonyl groups)), a tautomer prodrug or pharmaceutically acceptable salt of the compound or a solvate thereof, a thrombopoietin receptor activator, a preventive, therapeutic or improving agent for diseases against which activation of the thrombopoietin receptor is effective which contains the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient, and a platelet increasing agent containing the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient. It further relates to a pyrazolone compound represented by the formula (3)



Formula (3)

wherein  $R^7$  is a  $C_{2-14}$  aryl group (the  $C_{2-14}$  aryl group may be optionally substituted with one or more  $C_{1-6}$  alkyl groups, one or more  $C_{1-3}$  alkyl groups substituted with one or more fluorine atoms, one or more halogen atoms, one or more nitro groups, one or more  $C_{1-6}$  alkylcarbonyl groups, one or more hydroxyl groups or one or more amino groups (the hydroxyl group and the amino group may be substituted with a  $C_{1-6}$  alkyl group or a  $C_{1-6}$  alkylcarbonyl group)),  $R^8$  is a hydrogen atom, a  $C_{1-6}$  alkyl group, a  $C_{1-3}$  alkyl group substituted with one or more fluorine atoms or a  $C_{2-14}$  aryl group,  $R^9$  is a hydrogen atom, a  $C_{1-6}$  alkyl group, a  $C_{1-3}$  alkyl group substituted with one or more fluorine atoms or a  $C_{2-14}$  aryl group, and  $R^{10}$  is a  $C_{2-14}$  aryl group (the  $C_{2-14}$  aryl group is optionally substituted with one or more carboxyl groups, one or more sulfonic acid groups, one or more phosphonic acid groups, one or more ~~carbamido~~ ~~carbameoyl~~ groups, one or more ~~sulfamido~~ ~~sulfameoyl~~ groups, one or more ~~hydroxycarbamido~~ ~~hydroxycarbameoyl~~ groups, one or more ~~hydroxysulfamido~~ ~~hydroxysulfameoyl~~ groups, one or more tetrazole groups, one or more  $C_{1-6}$  alkoxy carbonyl groups or  $X(CYZ)_nCO_2H$  (wherein X is  $CH_2$ , O, S or  $NR^{11}$  ( $R^{11}$  is a hydrogen atom, a  $C_{1-6}$  alkyl group, a formyl group or a  $C_{1-6}$  alkylcarbonyl group), Y and Z are independently hydrogen atoms or  $C_{1-3}$  alkyl groups, and n is 0, 1, 2 or 3)), a tautomer, prodrug or pharmaceutically acceptable salt of the compound or a solvate thereof, a thrombopoietin receptor activator, a preventive, therapeutic or improving agent for diseases against which activation of the thrombopoietin receptor is effective which contains the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient, and a platelet increasing agent containing the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof

as an active ingredient. It still further relates to a pyrazolone compound represented by the formula (4)



Formula (4)

wherein  $R^{12}$  is a  $C_{2-14}$  aryl group (the  $C_{2-14}$  aryl group may be optionally substituted with one or more  $C_{1-6}$  alkyl groups, one or more  $C_{1-3}$  alkyl groups substituted with one or more fluorine atoms, one or more halogen atoms, one or more nitro groups, one or more  $C_{1-6}$  alkylcarbonyl groups, one or more hydroxyl groups or one or more amino groups (the hydroxyl group and the amino group may be substituted with a  $C_{1-6}$  alkyl group or a  $C_{1-6}$  alkylcarbonyl group)),  $R^{13}$  is a hydrogen atom, a  $C_{1-6}$  alkyl group, a  $C_{1-3}$  alkyl group substituted with one or more fluorine atoms or a  $C_{2-14}$  aryl group,  $R^{14}$  is a hydrogen atom, a  $C_{1-6}$  alkyl group, a  $C_{1-3}$  alkyl group substituted with one or more fluorine atoms or a  $C_{2-14}$  aryl group, and  $R^{15}$  is a  $C_{2-14}$  aryl group (the  $C_{2-14}$  aryl group is substituted with a substituent selected from a hydroxyl group, an amino group, a nitro group, a halogen atom, a cyano group, a  $C_{1-3}$  alkyl group substituted with one or more fluorine atoms, a carbamido ~~carbamo~~yl group and a sulfamido ~~sulfamo~~yl group (the carbamido ~~carbamo~~yl group and the sulfamido ~~sulfamo~~yl group may be substituted with a  $C_{1-6}$  alkyl group) and with a substituent selected from a carboxyl group, a sulfonic acid group, a phosphonic acid group, a carbamido ~~carbamo~~yl group, a sulfamido ~~sulfamo~~yl group, a hydroxycarbamido ~~hydroxycarbamo~~yl

group, a hydroxysulfamido ~~hydroxysulfamoyl~~ group, a tetrazole group, a C<sub>1-6</sub> alkoxy carbonyl group and X(CYZ)<sub>n</sub>CO<sub>2</sub>H (wherein X is CH<sub>2</sub>, O, S or NR<sup>16</sup> (R<sup>16</sup> is a hydrogen atom, a C<sub>1-6</sub> alkyl group, a formyl group or a C<sub>1-6</sub> alkyl carbonyl group), Y and Z are independently hydrogen atoms or C<sub>1-3</sub> alkyl groups, and n is 0, 1, 2 or 3)), a tautomer, prodrug or pharmaceutically acceptable salt of the compound or a solvate thereof, a thrombopoietin receptor activator, a preventive, therapeutic or improving agent for diseases against which activation of the thrombopoietin receptor is effective which contains the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient, and a platelet increasing agent containing the thrombopoietin receptor activator, a tautomer, prodrug or pharmaceutically acceptable salt of the thrombopoietin receptor activator or a solvate thereof as an active ingredient.

Please amend the paragraph beginning on page 25, line 11 as follows:

Substituents: a carboxyl group, sulfonic acid group, a phosphonic acid group, a carbamido ~~carbameoyl~~ group, a sulfamide group, a hydroxycarbamido ~~hydroxycarbameoyl~~ group, a hydroxysulfamido ~~hydroxysulfameoyl~~ group, CH<sub>2</sub>CO<sub>2</sub>H, OCH<sub>2</sub>CO<sub>2</sub>H, NHCH<sub>2</sub>CO<sub>2</sub>H, CH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>H and a tetrazole group.

Please amend the paragraph beginning on page 25, line 24 as follows:

Substituents: a carboxyl group, a sulfonic acid group, a phosphonic acid group, a carbamido ~~carbameoyl~~ group, a sulfamido ~~sulfameoyl~~ group, a hydroxycarbamido ~~hydroxycarbameoyl~~ group, a hydroxysulfamido ~~hydroxysulfameoyl~~ group, CH<sub>2</sub>CO<sub>2</sub>H, OCH<sub>2</sub>CO<sub>2</sub>H, NHCH<sub>2</sub>CO<sub>2</sub>H, CH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>H and a tetrazole group.

Please amend the paragraph beginning on page 26, line 17 as follows:

Substituent set A: a hydroxyl group, an amino group, a nitro group, a cyano group, a halogen atom, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms, a carbamido ~~earbamoyl~~ group and a sulfamido ~~sulfameoyl~~ group (the carbamido ~~earbamoyl~~ group and the sulfamido ~~sulfameoyl~~ group may be substituted with a C<sub>1-6</sub> alkyl group).

Please amend the paragraph beginning on page 26, line 23 as follows:

Substituent set B: a carboxyl group, a sulfonic acid group, a phosphonic acid group, a carbamido ~~earbamoyl~~ group, a sulfamido ~~sulfameoyl~~ group, a hydroxycarbamido ~~hydroxycarbameoyl~~ group, a hydroxysulfamido ~~hydroxysulfameoyl~~ group, CH<sub>2</sub>CO<sub>2</sub>H, OCH<sub>2</sub>CO<sub>2</sub>H, NHCH<sub>2</sub>CO<sub>2</sub>H, CH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>H and a tetrazole group.

Please amend the paragraph beginning on page 27, line 10 as follows:

Substituent set A: a hydroxyl group, an amino group, a nitro group, a cyano group, a halogen atom, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms, a carbamido ~~earbamoyl~~ group and a sulfamido ~~sulfameoyl~~ group (the carbamido ~~earbamoyl~~ group and the sulfamido ~~sulfameoyl~~ group may be substituted with a C<sub>1-6</sub> alkyl group).

Please amend the paragraph beginning on page 27, line 16 as follows:

Substituent set B: a carboxyl group, a sulfonic acid group, a phosphonic acid group, a carbamido ~~earbamoyl~~ group, a sulfamido ~~sulfameoyl~~ group, a hydroxycarbamido ~~hydroxycarbameoyl~~ group, a hydroxysulfamido ~~hydroxysulfameoyl~~ group, CH<sub>2</sub>CO<sub>2</sub>H, OCH<sub>2</sub>CO<sub>2</sub>H, NHCH<sub>2</sub>CO<sub>2</sub>H, CH<sub>2</sub>CH<sub>2</sub>CO<sub>2</sub>H and a tetrazole group.

Please amend the paragraph beginning on page 31, line 22 as follows:

19) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a  $C_{2-14}$  aryl group substituted with a carbamido ~~carbameoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 31, line 26 as follows:

20) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a phenyl group or pyridyl group substituted with a carbamido ~~carbameoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 32, line 4 as follows:

21) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a thienyl group, furyl group or pyridazinyl group substituted with a carbamido ~~carbameoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 32, line 9 as follows:

22) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a  $C_{2-14}$  aryl group substituted with a sulfamido ~~sulfameoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.



Please amend the paragraph beginning on page 32, line 13 as follows:

23) Pyrazolone compounds represented by the formula (3) wherein R<sup>10</sup> is a phenyl group or pyridyl group substituted with a sulfamido ~~sulfamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 32, line 18 as follows:

24) Pyrazolone compounds represented by the formula (3) wherein R<sup>10</sup> is a thienyl group, furyl group or pyridazinyl group substituted with a sulfamido ~~sulfamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 32, line 23 as follows:

25) Pyrazolone compounds represented by the formula (3) wherein R<sup>10</sup> is a C<sub>2-14</sub> aryl group substituted with a hydroxycarbamido ~~hydroxycarbamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 33, line 1 as follows:

26) Pyrazolone compounds represented by the formula (3) wherein R<sup>10</sup> is a phenyl group or pyridyl group substituted with a hydroxycarbamido ~~hydroxycarbamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 33, line 6 as follows:

27) Pyrazolone compounds represented by the formula (3) wherein R<sup>10</sup> is a thienyl group, furyl group or pyridazinyl group substituted with a hydroxycarbamido ~~hydroxycarbamoyl~~

group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 33, line 11 as follows:

28) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a  $C_{2-14}$  aryl group substituted with a hydroxysulfamido ~~hydroxysulfamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 33, line 16 as follows:

29) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a phenyl group or pyridyl group substituted with a hydroxysulfamido ~~hydroxysulfamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 33, line 21 as follows:

30) Pyrazolone compounds represented by the formula (3) wherein  $R^{10}$  is a thienyl group, furyl group or pyridazinyl group substituted with a hydroxysulfamido ~~hydroxysulfamoyl~~ group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 36, line 13 as follows:

40) Pyrazolone compounds represented by the formula (4) wherein  $R^{15}$  is a  $C_{2-14}$  aryl group substituted with a substituent selected from a nitro group, a cyano group, a  $C_{1-3}$  alkyl group substituted with one or more fluorine atoms, a carbamido ~~carbameoyl~~ group and a sulfamido ~~sulfameoyl~~ group (the carbamido ~~carbameoyl~~ group and the sulfamido ~~sulfameoyl~~ group may be

substituted with a C<sub>1-6</sub> alkyl group) and a halogen atom and with a carboxyl group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 36, line 23 as follows:

41) Pyrazolone compounds represented by the formula (4) wherein R<sup>15</sup> is a phenyl or pyridyl group substituted with a substituent selected from a nitro group, a cyano group, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms, a carbamido ~~carbameoyl~~ group and a sulfamido ~~sulfameoyl~~ group (the carbamido ~~carbameoyl~~ group and the sulfamido ~~sulfameoyl~~ group may be substituted with a C<sub>1-6</sub> alkyl group) and a halogen atom and with a carboxyl group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.

Please amend the paragraph beginning on page 37, line 6 as follows:

42) Pyrazolone compounds represented by the formula (4) wherein R<sup>15</sup> is a thienyl group, furyl group or pyridazinyl group substituted with a substituent selected from a nitro group, a cyano group, a C<sub>1-3</sub> alkyl group substituted with one or more fluorine atoms, a carbamido ~~carbameoyl~~ group and a sulfamido ~~sulfameoyl~~ group (the carbamido ~~carbameoyl~~ group and the sulfamido ~~sulfameoyl~~ group may be substituted with a C<sub>1-6</sub> alkyl group) and a halogen atom and with a carboxyl group, tautomers, prodrugs or pharmaceutically acceptable salts of the compounds or solvates thereof.